1616 18 PATENTS

## IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In a Application of:

· , ) ) RECEIVED

DEC 0 5 2003

Serial No. 10/077,142

Art Unit: **1616** 

**TECH CENTER 1600/2900** 

201141 1 (0. 10/01/1,11

'Amato et al.

Examiner: Badio, Barbara P.

Filed: **February 15, 2002** 

For: Estrogenic Compounds as Anti-Mitotice Agents

## SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT

Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

Sir:

The citation of information on the attached Form PTO/SB/08 is made pursuant to 37 C.F.R. §§ 1.56, 1.97, and 1.98. A copy of each cited item is enclosed.

As this Information Disclosure Statement is being filed after the issuance of an office action on the merits, pursuant to 37 C.F.R. §1.97(c)(2), the fee of \$180, as set forth in §1.17(p), is enclosed.

The citation of this information does not constitute an admission of priority or that any cited item is available as a reference, or a waiver of any right the applicant may have under applicable statutes, Rules of Practice in patent cases, or otherwise.

Respectfully submitted,

KILPATRICK STOCKTON LLP Suite 2800, 1100 Peachtree Street

(404) 815-6500

Our Docket:

05213-3000 (43170-269288)

Sima Singadia Kulkarni Reg. No. 43,732

I hereby certify that this correspondence is being deposited with the United States Postal Service as first class mail in an envelope addressed to: Commissioner for Patents, P.O. Box 1450, Alexandria VA 22313-1450, on November 25, 2003.

Sima Singadia Kalkarni – Reg. No: 43,73

Approved for use through 10/31/2002 OMB 0651-0031

U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE Under the Paperwork Reduction Act of 1995 required to respond to a collection of information unless it contains a valid OMB control number TECH OF STONS TO STONE OF STON Complete if Known Substitute for Form 1449/A/PTO Application No. 10/077,142 Filing Date 02/15/2002 INFORMATION DES First Named Inventor D'Amato et al. STATEMENT BY APPLIES Group Art Unit 1616 (use as many sheets as necessary) **Examiner Name** Badio, Barbara P. Sheet 1 of 15 Attorney Docket Number 05213-3000 (43170-269288) U.S. PATENT DOCUMENTS U.S. Patent Document Date of Publication Examiner Cite Pages, Columns, Lines, Where Kind Code Initials Name of Patentee or of Cited Document No.1 Relevant Passages or Number<sup>4</sup> (if known) MM-DD-YYYY Applicant of Cited Document Relevant Figures Appear 1 3,166,577 Ringold et al. 01/19/1965 2 3,562,260 De Ruggieri et al. 02/09/1971 3 4,552,758 Murphy et al. 11/12/1985

02/28/1989

Leibovich et al.

4

4,808,402

1	Examiner	Date	
	Cionatura	Date	
1	Signature	Considered	<u> </u>
- 1	L	00110100100	

Approved for use through 10/31/2002 OMB 0651-0031

U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE Under the Paperwork Reduction Act of 1995 required to respond to a collection of information unless it contains a valid OMB control number Complete if Known TOENTER STOOLSON Application No. 10/077,142 Substitute for Form 1449/A/PTO Filing Date 02/15/2002 INFORMATION DES First Named Inventor D'Amato et al. STATEMENT BY A 1616 Group Art Unit (use as many sheets as necessary **Examiner Name** Badio, Barbara P. 05213-3000 (43170-269288) Sheet 2 of 15 Attomey Docket Number FOREIGN PATENT DOCUMENTS Foreign Patent Document Date of Publication Pages, Columns, Lines, Where Examiner Cite Kind Code of Cited Document Relevant Passages or Name of Patentee or Initials No.1 Office3 MM-DD-YYYY Number<sup>4</sup> Relevant Figures Appear (if known) Applicant of Cited Document 5 wo 88/08002 A1 The Upjohn Company 10/20/1988 wo Αl 93/19746 ORION-YH-TYMA OY 10/14/1993

Examiner		Date	
Ciamatura		Date	
Signature	• [	Considered	

Approved for use through 10/31/2002 OMB 0651-0031 U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE

nopperions are required to respond to a collection of information unless it contains a valid OMB control number Under the Paperwork Reduction Act of 15 Complete if Known 10/077,142 Application No. Substitute for Form 1449/A/PTO Filing Date 02/15/2002 INFORMATION First Named Inventor D'Amato et al. STATEMENT BY Group Art Unit 1616 (use as many sheets as necessary) Badio, Barbara P. **Examiner Name** of 05213-3000 (43170-269288) Sheet 3 15 Attorney Docket Number OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the Examiner Cite item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), **Initials** No. publisher, city and/or country where published Author: Title: Publ: Research Plus Catalog Pages: 50-58 Vol/Iss: Date: 1993 Title: Registry Number: 56933-77-8 8 Author: Publ: Chemical Abstracts Pages: Date: Vol/iss: Title: Registry Number: 56933-78-9 Author: Publ: Chemical Abstracts Pages: Vol/lss: Date: Author: Title: Registry Number: 57380-15-1 10 Chemical Abstracts Pages: Vol/lss: Date: Title: Registry Number: 71782-94-0 11 Author: Publ: Chemical Abstracts Pages: Vol/lss: Date: Title: Registry Number: 71782-95-1 Author: 12 Publ: Chemical Abstracts Vol/Iss: Pages: Date: Title: Registry Number: 101277-11-6 Author: 13 Publ: Chemical Abstracts Pages: Vol/Iss: Date: Date Examiner Considered Signature

'Unique citation designation number. 'See attached Kinds of U.S. Patent Documents. 'Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). 'For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. 'Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. 'Applicant is to place a check mark here if English language translation is attached.

Approved for use through 10/31/2002 OMB 0651-0031

U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE Under the Paperwork Reduction Act of 199 are required to respond to a collection of information unless it contains a valid OMB control number 2003 2003 CONTRACTION 2000 Complete if Known Application No. Substitute for Form 1449/A/PTO 10/077,142 Filing Date 02/15/2002 INFORMATION (B) First Named Inventor D'Amato et al. STATEMENT BY Group Art Unit 1616 (use as many sheets as necessary **Examiner Name** Badio, Barbara P. Sheet of 15 Attorney Docket Number 05213-3000 (43170-269288) OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the Examiner Cite T٠ item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), Initials No. publisher, city and/or country where published Author: Title: Registry Number: 101429-40-7 Publ: Chemical Abstracts Vol/Iss: Pages: Date: Author: 15 Title: Registry Number: 162853-20-5 Publ: Chemical Abstracts Pages: Vol/Iss: Date: 16 Author: ABOULWAFA et al. Title: Synthesis and evaluation for uterotrophic and antiimplantation activities of 2-substituted estradiol derivatives Publ: Steroids Vol/Iss: 57 Pages: 199-204 Date: 04/1992 Title: Vascular reactions of normal and malignant tumors in vivo. I. Vascular reactions of Author: ALGIRE, G.H. et al. mice to wounds and to normal and neoplastic transplants Journal of the National Cancer Institute Vol/lss: 6 Pages: 73-85 Date: 08/1945 Author: ALIEV et al. Title: 54929q Synthesis of cycloalkyl derivatives of dihydric phenols and their ethers 18 Chemical Abstracts Vol/Iss: 72 Pages: 370 Date: 1970 Title: Tumor Necrosis Factor Production in Patients with Leprosy 19 Author: BARNES et al. Infection and Immunity Vol/lss: 60 (4) Pages: 1441-1446 Date: 04/1992 Author: BINDRA et al. 20 Title: Studies in Antifertility Agents. 8. Seco Steroids. 2. 5,6-Secoestradiol and Some Related Compounds Journal of Medicinal Chemistry Vol/lss: 18 (9) Pages: 921-925 Date: 1975

Examiner	Date	]
Signature	Considered	

Approved for use through 10/31/2002 OMB 0651-0031 U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE

are required to respond to a collection of information unless it contains a valid OMB control number Under the Paperwork Reduction Act of 1995 Complete if Known Substitute for Form 1449/A/PTO Application No. 10/077,142 DEC 0 1 2003 Filing Date 02/15/2002 **INFORMATION** SIS First Named Inventor D'Amato et al. STATEMENT BY Group Art Unit 1616 (use as many sheets as necessary) **Examiner Name** Badio, Barbara P. Sheet of 5 15 Attorney Docket Number 05213-3000 (43170-269288) OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the Examiner Cite item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), T٠ Initials No. publisher, city and/or country where published Author: BLICKENSTAFF et al. 21 Title: SYNTHESIS OF SOME ANALOGS OF ESTRADIOL Publ: Steroids 5 4 1 Vol/Iss: 46 (4,5) Pages: 889-902 Date: 10/1985 Author: BOYCE et al. Title: Some Preliminary Synthetical Studies with 5,6,7,8-Tetra-hydro-8-methylindane -1,5-dione Publ: <u>Jnknown</u> Vol/iss: Pages: 4547-4553 Date: 1960 Author: BRANDI et al. Title: Bone endothelial cells as estrogen targets (Abstract only) 23 Publ: Calcif. Tissue Int. Vol/Iss: 53 (5) Pages: 312-317 Date: 1993 Title: Interstitial chemotherapy with drug polymer implants for the treatment of recurrent Author: BREM, H. et al. gliomas Journal of Neurosurgery Vol/iss: 74 Pages: 441-446 Date: 03/01/91 25 Author: CAMBIE et al. Title: Aromatic Steroids. Part I. Oxidation Products of 3-Methoxyestra-1,3,5(10)-triene- 17βyl Acetate Publ: . Chem. Soc. Vol/Iss: Pages: 2603-2608 Date: 26 Author: COREY et al. Title: Applications of N,N-Dimethylhydrazones to Synthesis. Use in Efficient, Positionally and Stereochemically Selective C-C Bond Formation; Oxidative Hydrolysis to Publ: Carbonyl Compounds Tetrahedron Letters Vol/Iss: I Pages: 3-6 Date: 1976 Title: Facile Conversion of N,N-Dimethylhydrazones to Cabonyl Compounds by Cupric Author: COREY et al. Ion-Catalyzed Hydrolysis Publ: Tetrahedron Letters Vol/Iss:41 Pages: 36678-3668 Date: 1976

Examiner	Date	· · · · ·
Signature	Considered	

PTO/SB/08A (08-00) Approved for use through 10/31/2002 OMB 0651-0031 U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE ms are required to respond to a collection of information unless it contains a valid OMB companymber Under the Paperwork Reduction Act of Complete if Known Application No. 10/077,142 Substitute for Form 1449/A/PTO 02/15/2002 Filing Date **INFORMATION** First Named Inventor D'Amato et al. STATEMENT BY & BI Group Art Unit 1616 (use as many sheets as necessary) **Examiner Name** Badio, Barbara P. Sheet 6 of 15 Attorney Docket Number 05213-3000 (43170-269288) OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the Examiner Cite ۲ item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), Initials No.1 publisher, city and/or country where published Title: Cotton effect of the styrene chromophore (Abstract only) Author: CRABBE, P. 28 Publ: Chem. Ind. Vol/Iss:27 Pages: 917-918 Date: 1969 Title: Seco-Oestradiols and Some Non-Steroidal Oestsrogens: Structural Correlates of 29 Author: DURANI et al. Oestrogenic Action Publ: Journal of Steroid Biochemistry

•		1				
	<u></u>		Vol/lss: 11	Pages: 67-77	Date:	1979
	30	Author: EDER et al.	Title: Synthese von Ostradiol (in	German - No translation avail	lable)	
		Publ: Chem. Ber.				
	<u></u>		Vol/lss: 109	Pages: 2948-2953	Date:	1976
	31	Author: EMONS et al. Publ: FOCUS MHL	Title: Modulation der hypophysa Ostrogene	ren Sekretion von Luteinisiere	endem Hormo	n (LH) durch
			Vol/lss:3	Pages: 221-228	Date:	1986
• •	32	Author: FANCHENKO et al.	Title: Characterisitics of the guin	ea nig uterus estrogen recentor	r system (Abs	tract only)
		Publ: Byull. Eksp. Biol. Med.		on pig moras sonogen receptor		,
			Vol/lss: 85 (4)	Pages: 467-470	Date:	1978
	33			Pages: 467-470	Date:	1978
	33	Byull. Eksp. Biol. Med.  Author: FEVIG et al.  Publ:	Vol/lss: 85 (4)	Pages: 467-470	Date:	1978
	33	Byull. Eksp. Biol. Med.  Author: FEVIG et al.  Publ:	Vol/Iss: 85 (4) Title:A Short, Stereoselective	Pages: 467-470  Route to 16α(Substituted-a  Pages: 247-251	Date:  kyl)estradiol	1978 I Derivatives
		Byull. Eksp. Biol. Med.  Author: FEVIG et al.  Publ:  Journal of Organic Chemistry	Vol/Iss: 85 (4)  Title: A Short, Stereoselective  Vol/Iss: 52	Pages: 467-470  Route to 16α(Substituted-a  Pages: 247-251	Date:  kyl)estradiol	1978 I Derivatives

Examiner Date Considered Considered

'Unique citation designation number. 'See attached Kinds of U.S. Patent Documents. 'Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). 'For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. 'Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. 'Applicant is to place a check mark here if English language translation is attached.

Approved for use through 10/31/2002 OMB 0651-0031

U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE ans are required to respond to a collection of information unless it contains a valid OMB control number Under the Paperwork Reduction Act of 19 Complete if Known 10/077,142 Application No. Substitute for Form 1449/A/PTO Filing Date 02/15/2002 INFORMATIO First Named Inventor D'Amato et al. STATEMENT BY Group Art Unit 1616 (use as many sheets as **Examiner Name** Badio, Barbara P. 05213-3000 (43170-269288) Sheet of 7 15 Attorney Docket Number OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the Examiner Cite ť item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), Initials No.1 publisher, city and/or country where published Author: FIESER et al. Title: N-Methylformanilide 35 Publ: Organic Synthesis Collective Volume 3 Vol/Iss:3 Pages: 590-591 Date: 1955 Author: FLOHE et al. Title: Studies on the Hypothetical Relationship of Thalidomide-induced Embryopathy and 36 Collagen Biosynthesis Arzneimitte/Forschung (Germany West) Vol/lss: 31 (2) Pages: 315-320 Date: 01/01/81 Title: Angiogenesis Inhibition and Tumor Regression Caused by Heparin or a Heparin Author: FOLKMAN et al. Fragment in the Presence of Cortisone Publ: Science 5 4 1 Vol/lss: 221 Pages: 719-725 Date: 08/19/83 Title: Tumor Angiogenesis: Therapeutic Implications Author: FOLKMAN, J. Publ: New England Journal of Medicine Vol/Iss: 285 (21) Pages: 1182-1186 Date: 11/18/71 Title:Induction of Angiogenesis During the Transition from Hyperplasia to Neoplasia Author: FOLKMAN, J. et al. Publ: Nature Vol/lss: 339 Pages: 58-61 Date: 05/04/89 40 Author: FOLKMAN, J. et al. Title: Tumor Behavior in Isolated Perfused Organs in vitro Growth and Metastases of Biopsy Material in Rabbit Thyroid and Canine Intestinal Segment Publ: Annals of Surgery Pages: 491-502 Vol/lss: 164(3) Date: 09/01/66 Title: Mannich reaction of estrone 41 Author: GANDHI et al. Publ: Journal of Indian Chem. Soc. Vol/Iss: 39 Pages: 306-308 Date: 1962

Examiner	Date	
Signature	Considered	

Approved for use through 10/31/2002 OMB 0651-0031 U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE

Under the Paperwork Reduction Act of 199 to proper are required to respond to a collection of information unless it contains a valid OMB control number Complete if Known Application No. 10/077,142 Substitute for Form 1449/A/PTO DEC 0 1 2003 Filing Date 02/15/2002 **INFORMATION DISCLOS** First Named Inventor D'Amato et al. STATEMENT BY APPLE Group Art Unit 1616 (use as many sheets as necessary) **Examiner Name** Badio, Barbara P. 05213-3000 (43170-269288) Sheet 8 of 15 Attorney Docket Number OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the Examiner Cite T٤ item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), Initials No.1 publisher, city and/or country where published Author: GASLINI et al. Title: Reaction of Eugenol with Synthesis Gas. Synthesis of 5,6,7,8-Tetrahydro 42 -3-methoxy-2-napthol Journal of Organic Chemistry Vol/lss: 29 (5) Pages: 1177-1180 Date: 05/1964 43 Author: GIMBRONE, M.A. et al. Title: Tumor Growth and Neovascularization: An Experimental Model Using the Rabbit Cornea Publ: Journal of the National Cancer Institute Vol/Iss: 52(2) Pages: 413-427 Date: 02/74 Author: GIMBRONE, M.A. et al. Title: Tumor dormancy in vivo by Prevention of Neovascularization Publ: Journal of Experimental Medicine Vol/iss: 136 Pages: 261-276 Date: 1972 Author: GONZALEZ et al. Title: Synthesis and Pharmacological Evaluation of 8œEstradiol Derivatives Publ: Steroids Vol/Iss: 40 (2) Pages: 171-187 Date: 9/1982 Author: GROSS, J.L. et al. Title: Modulation of Solid Tumor Growth in vivo by bFGF (Abstract only) Proceedings of the American Association of Cancer Research Vol/lss:31 Pages: 79 Date: 03/90 47 Author: GUNZLER, V. Title: Thalidomide-A Therapy for the Immunological Consequences of HIV Infection? Publ: Medical Hypothesis Vol/lss: 30 (2) Pages: 105-109 Date: 10/89 Author: GUPTA et al. Title: Antifertility Agents. XIV. Secosteroids. VII. Synthesis of 2a and 2B, 6β-dimethyl- 3β-(p-hyroxyphenyl)-trans-bicyclo[4.3.0]nonan-7-ones and some Publ: related compounds (Abstract only) Indian Journal of Chemistry Vol/Iss: 13 (7) Pages: 759-760 Date: 1975

Examiner Signature		Date Considered	
Hilminus siasaine	defends and a 20 months of the CVI of December 177.	000 11 11 1	

U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE

Under the Paperwork Reduction Act of 199 are required to respond to a collection of information unless it contains a valid OMB control number Complete if Known Substitute for Form 1449/A/PTO Application No. 10/077,142 Filing Date 02/15/2002 **INFORMATION DISCL** First Named Inventor D'Amato et al. STATEMENT BY Group Art Unit 1616 (use as many sheets as necessary **Examiner Name** Badio, Barbara P. Sheet 9 of 15 Attorney Docket Number 05213-3000 (43170-269288 OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the Examiner Cite item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), Initials No.1 publisher, city and/or country where published Author: GUPTA et al. Title: Studies in Antifertility Agents. Part XVIII. 2α6β-Diethyl-3β-(p-hydroxyphenyl) 49 -trans-bicyclo[4.3.0]nonan-7β-ol and 6 $\beta$ -methyl-3 $\beta$ -(p-hydroxyphenyl)-2 $\alpha$ -propyl- trans-bicyclo[4.3.0]nonan-7 $\beta$ ol Indian Journal of Chemistry (Abstract only) Vol/lss: 19B (10) Pages: 886-890 Author: GUTIERREZ-RODRIGUEZ Title: Treatment of Refractory Rheumatoid Arthritis - The Thalidomide Experience 50 Publ: The Journal of Rheumatology Vol/lss: 16 (2) Pages: 158-163 Date: 02/89 Author: GUTIERREZ-RODRIGUEZ, Title: Thalidomide - A Promising New Treatment for Rheumatoid Arthritis Publ: Arthritis and Rheumatism Vol/lss: 27 (10) Pages: 1118-1121 Date: 10/84 Title: THE SPECIFICITY OF THE ESTROGEN RECEPTOR OF HUMAN UTERUS Author: HAHNEL et al. Publ: Journal of Steroid Biochemistry Vol/lss: 4 Pages: 21-31 Date: 1973 Author: HANDLEY et al. Title: Chronic bullous disease of childhood and ulcerative colitis British Journal of Dermatology Vol/lss: 127 (40) Pages: 67-68 Date: 07/01/92 54 Author: HENEY et al. Title: Thalidomide treatment for chronic graft-versus-host disease Publ: British Journal of Haematology Vol/lss: 78 (1) Pages: 23-27 Date: 05/91 Author: HOLKER et al. Title: The Reactions of Estrogens with Benzeneseleninic Anhydride and Hexamethyldisilazane Publ: . Chem. Soc. Perkin Trans. Vol/lss: I Pages: 1915-1918 Date: 1982 Examiner

Examiner
Signature

Date
Considered

'Unique citation designation number. 'See attached Kinds of U.S. Patent Documents. 'Enter Office that issued the document, by the two-letter code

(WIPO Standard ST.3). 'For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. 'Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. 'Applicant is to place a check mark here if English language translation is attached.

U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE

Under	the Pag	erwork Reduction Act of 1995, no perso	are required to re	espond to a collection	n of information u	nless it contains a	valid OMB cor	trol nun
		70	8			if Known	/	O.
Substitut	e for Fo	rm 1449/A/PTO	1 2003 5	Application No	0.	10/077,142	•	K)
,	<b>3</b> 1 <b>2</b> 2	DEC 0	<b></b>	Filing Date		02/15/2002	80, Q	<del>′,C</del>
		DRMATION DISCLOS		First Named In	nventor	D'Amato et al.	Ca	<u>~</u>
	STA	TEMENT BY APPLIE	T	Group Art Uni	it	1616	1/2	5
		(use as many sheets as necessary)		Examiner Nan	ne	Badio, Barbara	P. **	-
Shee	t	10 of 15		Attomey Dock	et Number	05213-3000 (43		0
				Taloniey Boar	et Humber	03213-3000 (43	170-209200)	-3
_	OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS							
Examiner	Cite	Include name of the auth	or (in CAPITAL L	ETTERS), title of th	e article (when app	propriate), title of	the	T
Initials	No.'	item (book, magazine, jou	publisher, city a	nd/or country where	published	ume-issue number	(s),	
	56	Author: HORL, A. et al.	Title: Suppression	of Solid Tumor Gro	owth by Immunone	cutralizing Monocl	onal Antibody	+
		Publ:	Against Hur	nan Basic Fibroblas	ts Growth Factor	g		ł
		Cancer Research						
			Vol/lss:5	1	Pages: 6180-61	84 Date:	44/45/04	
	57	Author: IMAMURA et al.					11/15/91	<u> </u>
	3/		a lue: Meulod for	Manufacture of Dihy	dric Phenois (Abs	tract only)		
		Publ: USPATFULL 76:20259 US 3,950,437						
		<u> </u>			<b>D</b>			
	<u> </u>		Vol/lss:		Pages:	Date:	04/13/1976	
	58	Author: INGBER, D. et al.	Title:Synthetic an	alogues of fumagilli	n that inhibit angio	ogenesis and suppr	ess tumor grow	rth .
		Publ:						
		Nature						
			Vol/Iss: 34	18	Pages: 555-557	Date:	12/06/90	Ш
	59	Author: IRIARTE et al.	Title:Steroids (XC	CIV). Synthesis of 2-	methyl and 1,2-dir	nethyl estrogens (A	Abstract only)	abla
		Publ:						1 1
		<u>Tetrahedron</u>						
			Vol/lss:3		Pages: 28-36	Date:	1958	
	60	Author: JHINGRAN et al.	Title: Studies in A	Intifertility Agents -	Part XLI: Secosi	teroids-x: Synthe	ses of Various	
		  Publ:	Stereoisom	ers of (+-)-2,6β-die	thyl-7α-ethynyl-3	-(p-hydroxyphen	yl)-trans-	
		Steroids	Dicyclo [4.3.	.0)nonan-7β-ol.				1 1
			Vol/Iss: 42	(6)	Pages: 627-634	Date:	1983	
T I	61	Author: KARWAT		nd Recovery of Hydr				+
	01		i ide. Separadon ai	id Recovery of Hydr	logen Sulfide from	Hydrocarbon Mix	ture	
		Publ: Caplus DE 1103310						
ļ		<u>Gapius 515 1105510</u>	M-10.					
1			Vol/Iss:		Pages:	Date:	09/02/1959	ل
ŀ	62	Author: KIM, K.J. et al.	Title:Inhibition of Y Tumor Grown	Vascular Endothelial	Growth Factor-in	duced Angiogenes	is Suppresses	
l		Publ:	Tutioi Olow	MI 111 A 140				
		<u>Nature</u>						
		·	Vol/lss: 362	2	Pages: 841-844	Date:	04/29/93	Ш
							_	
	_							
						<del></del>		

Examiner	Date	
Signature	Considered	

Approved for use through 10/31/2002 OMB 0651-0031 U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE persons are required to respond to a collection of information unless it contains a valid OMB control number Under the Paperwork Reduction Act of The Contraction of the Contracti Complete if Known Substitute for Form 1449/A/PTO Application No. 10/077.142 Filing Date 02/15/2002 INFORMATION: DISCLO First Named Inventor D'Amato et al. STATEMENT BY XPRE Group Art Unit 1616 (use as many sheets as necessary) **Examiner Name** Badio, Barbara P. Sheet of 11 15 Attorney Docket Number 05213-3000 (43170-269288) OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the Examiner Cite T٩ item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), **Initials** No.1 publisher, city and/or country where published Author: KNIGHTON, D. et al. Title: Avascular and Vascular Phases of Tumour Growth in the Chick Embyo Publ: British Journal of Cancer Vol/lss:35 Pages: 347-356 Date: 1977 Author: KOLE et al. 64 Title: Studies in Antifertility Agents. 11. Secosteroids. 5. Synthesis of 9,11-Secoestradiol Publ: Journal of Medicinal Chemistry Vol/lss: 18 (7) Pages: 765-766 Date: 1975 65 Author: KOVACS et al. Title: Steroids. XXIII. Synthesis of 2- and 4-hydroxy and 2,4-dihydroxy derivatives of estrone and estradiol (Abstract only) Publ: Acta Phys.Chem. Vol/lss: 19 (3) Pages: 287-290 Date: 1973 Author: LEWIS, Richard J. Title: Hawley's Condensed Chemical Dictionary Vol/Iss: Pages: 577 Date: 01/1993 Author: LEWIS, Richard J. Title: Hawley's Condensed Chemical Dictionary Vol/iss: Pages: 128-129 Date: 01/1993 68 Author: LIEN, W. et al. Title: The blood supply of experimental liver metastases. II. A Microcirculatory study of the normal and tumor vessels of the liver with the use of perfused silicone rubber Publ: Surgery Vol/lss: 68 (2) Pages: 334-340 08/1970 Author: LIU et al. Title: Total Synthesis of (+-) -Δ<sup>9(12)</sup> Capnellene Publ: Tetrahedron Letters Vol/lss: 26 (40) Pages: 4847-4850 Date: 1985

Examiner	Date	
Signature	Considered	

Approved for use through 10/31/2002 OMB 0651-0031

U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE
Under the Paperwork Reduction Act of 1990 no persons are required to respond to a collection of information unless it contains a valid OMB control number Complete if Known DEC 0 1 2003 Substitute for Form 1449/A/PTO Application No. 10/077,142 Filing Date 02/15/2002 INFORMATION DISC First Named Inventor D'Amato et al. STATEMENT BY APPLICAN Group Art Unit 1616 (use as many sheets as necessary) **Examiner Name** Badio, Barbara P. Sheet 12 of 15 Attorney Docket Number 05213-3000 (43170-269288) OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the Examiner Cite item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), T٤ Initials No.1 publisher, city and/or country where published Author: LOOZEN et al. Title: An approach to the synthesis of 7.beta.-amino estrogens (Abstract only) Recl.: J.R. Neth.Chem. Soc. Vol/lss: 102 (10) Pages: 433-437 Date: 1983 Author: MARO et al. Title: Mechanism of Polar Body Formation in the Mouse Oocyte: An Interaction Between the Chromosomes, the Cytoskeleton and the Plasma Membrane Publ: Journal of Embryology and Experimental Morphology Vol/lss:92 Pages: 11-32 Date: 1986 Title: Inhibition of synaptosomal high-affinity uptake of dopamine and serotonin by estroger Author: MICHEL et al. agonists and antagonists (Abstract only) Publ: Biochem. Pharmacol. Vol/Iss: 36 (19) Pages: 3175-3180 Date: 1987 Title: Steroids. L1. Aromatization reaction of the cross-conjugated dienone system by Zinc 73 Author: MORISAKI et al. 9. (Abstract only) Chem. Pharm. Bull. Vol/Iss: 14 (8) Pages: 866-872 Date: 1966 74 Author: NAAFS et al. Title: Thalidomide Therapy An Open Trial International Journal of Dermatology Vol/lss: 24 (2) Pages: 131-134 Date: 03/1985 Author: NAMBARA et al. 75 Title: Microbial transformation products derived from steriods. 1. Synthesis of 1,2- and 3-dimethoxy-4-methylestratrienes (Abstract only) Publ: Chem. Pharm. Bull. Vol/iss: 20 (2) Pages: 336-342 Date: 1972 Author: NAMBARA et al. Title: Synthesis of 16β-Oxygenated Catechol Estrogen Methyl Ethers, New and 76 **Potential Metabolites** Publ: Chemical & Pharmaceutical Bulletin Vol/lss: 23 (7) Pages: 1613-1616 Date: 07/1975

Examiner	Date	
Signature	Considered	

Approved for use through 10/31/2002 OMB 0651-0031

U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE no persons are Under the Paperwork Reduction Act of equired to respond to a collection of information unless it contains a valid OMB control number Complete if Known Substitute for Form 1449/A/PTO Application No. 10/077,142 Filing Date 02/15/2002 INFORMATION DE First Named Inventor D'Amato et al. STATEMENT BY APPLICA Group Art Unit 1616 (use as many sheets as necessary) **Examiner Name** Badio, Barbara P. Sheet of 13 15 Attorney Docket Number 05213-3000 (43170-269288) OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the Examiner Cite item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), Initials No. publisher, city and/or country where published Author: NEWKOME et al. Title: Synthesis of Simple Hydrazones of Carbonyl Compounds by an Exchange Reaction 77 Publ: Journal of Organic Chemisty Vol/lss:31 Pages: 677-681 Date: 03/1966 Title: Elevated Levels of the Angiogenic Peptide Basic Fibroblast Growth Factor in Urine of 78 Author: NGUYEN, M. et al. **Bladder Cancer Patients** Journal of the National Cancer Institute Vol/Iss: 85 (3) Pages: 241-242 Date: 02/03/93 Author: NUMAZAWA et al. Title: Novel and Regiospecific Synthesis of 2-Amino Estrogens via Zincke Nitration Publ: Steroids 5 4 1 Vol/lss: 41 (5) Pages: 675-682 Date: 1983 Author: PERT et al. Title: Preparations of 2,4-disubstituted estradiols (Abstract only) 80 Publ: Australian Journal of Chemistry Vol/Iss: 42 (3) Pages: 421-432 Date: 1989 Author: PETERS et al. Title: 17-Desoxy Estrogen Analogues Journal of Medicinal Chemistry Pages: 1642-1652 Vol/lss: 32 (7) Date: 82 Author: PFEIFFER et al. Title: Are catechol estrogens obligatory mediators of estrogen action in the central nervous system? I. Characterization of pharmacological probes with different receptor binding Publ: affinities and catechol estrogen formation rates (Abstract only) Journal of Endocrinology Vol/lss: 110 (3) Pages: 489-497 Date: 1986 Author: POWELL et al. Title: Investigation and Treatment of Orogenital Ulceration; studies on a Possible Mode of Action of Thalidomide Publ: British Journal of Dermatology

Examiner	Date	
Signature	Considered	

Vol/Iss: 113 Supp. 28

Pages: 141-144

Date:

07/1985

Approved for use through 10/31/2002 OMB 0651-0031 U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE

B contraction on the state of t Under the Paperwork Reduction Act of 19 equired to respond to a collection of information unless it contains a valid OMB control Complete if Known Substitute for Form 1449/A/PTO Application No. 10/077,142 Filing Date 02/15/2002 INFORMATION DIS First Named Inventor D'Amato et al. STATEMENT BY APPE Group Art Unit 1616 (use as many sheets as necessary) **Examiner Name** Badio, Barbara P. Sheet of Attorney Docket Number 14 15 05213-3000 (43170-269288) OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the Examiner Cite item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), Initials No.1 publisher, city and/or country where published Author: SHISHKINA et al. Title: Synthesis and properties of condensed heterocyclic derivatives of estra-4, 9-dien-17.beta.-ol-3-one (Abstract only) Publ: Khim.-Farm. Zh. Vol/lss: 8 (1) Pages: 7-11 Date: 1974 Author: SIDKY et al. Title: Inhibition of Angiogenesis by Interferons: Effects on Tumor- and Lymphocyte-induced 85 Vascular Responses Publ: Cancer Research Vol/Iss: 47 Pages: 5155-5161 Date: 10/01/87 Author: SIRACUSA et al. Title: The effect of microtubule- and microfilament-disrupting drugs on preimplantation mouse embryos (Abstract only) Jourant of Embryology and Experimental Morphology Pages: 71-82 Vol/Iss: 60 Date: 12/1980 Author: SPYRIOUNIS et al. Title: Copper (II) complex of an estradiol derivative with potent antiinflammatory properties (Abstract only) Publ: Arch. Pharm. Vol/Iss: 324 (9) Pages: 533-536 Date: 1991 Author: SRIVASTAVA, A. et al. Title: The Prognostic Significance of Tumor Vascularity in Intermediate-Thickness (0.76-4.0 mm Thick) Skin Melanoma American Journal of Pathology Vol/Iss: 133 (2) Pages: 419-424 Date: 11/1988 89 Author: STARKOV et al. Title: Mono- and Dialkylation of Guaiacol by Olefins on KU-2 Cation Exchanger (Abstract only) Publ-Zhumal Prikladnoi Khimii Pages: 688-690 Vol/lss:41 (3) Date: 1968 Author: TAYLOR, S. et al. Title: Protamine is an Inhibitor of Angiogenesis Publ: <u>Nature</u> Vol/Iss: 297 Pages: 307-312 Date: 05/27/82

Examiner Signature		Date Considered	
12.2.2	defends to 20 m to twice our and a line of	000 11 11 11 1	

PTO/SB/08A (08-00) Approved for use through 10/31/2002 OMB 0651-0031 U.S. Patent and Trademark Office: U.S. DEPARTMENT OF MMERCE Under the Paperwork Reduction Act of 199 quired to respond to a collection of information unless it contains a valid OMB control number Complete if Known Substitute for Form 1449/A/PTO Application No. 10/077.142 Filing Date 02/15/2002 INFORMATION DIS First Named Inventor D'Amato et al. STATEMENT BY APPE Group Art Unit 1616 (use as many sheets as necessary) **Examiner Name** Badio, Barbara P. Sheet 15 of 15 Attorney Docket Number 05213-3000 (43170-269288) OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the Examiner Cite item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), Initials No.1 publisher, city and/or country where published Title: In Vitro Activity of Thalidomide Against Mycobacterium avium Complex Author: VICENTE et al. Publ: Archives of Internal Medicine Vol/lss: 153 (4) Pages: 534 Date: 02/22/93 Author: WANG et al. Title: Photoaffinity labeling of human placental estradiol 17.beta.-dehydrogenase with 2- and 4-azidoestrone, 2- and 4-azidoestradiol (Abstract only) Publ: Shengwu Huaxue Zazhi Vol/Iss: 8 (6) Pages: 715-718 Date: 1992 Title: Tumor angiogenesis: A New Significant and Independent Prognostic Indicator in Author: WEIDNER, N. et al. Early-Stage Breast Carcinoma Publ: Journal of the National Cancer Institute Vol/Iss: 84 Pages: 1875-1887 Date: 12/16/92 Author: WEIDNER, N. et al. Title: Tumor Angiogenesis Correlates with Metastasis in Invasive Prostate Carcinoma American Journal of Pathology Vol/Iss: 143 (2) Pages: 401-409 Date: 08/1993 95 Author: WEIDNER, N. et al. Title: Tumor Angiogenesis and Metastasis-Correlation in Invasive Breast Carcinoma New England Journal of Medicine Vol/Iss: 324 (1) Pages: 1-8 Date: 01/03/91 Author: WELSCH et al. 96 Title: Staphylostatic Activity of Some New Diphenols, Napthols, and Chalcones (Abstract only) Publ: Experientia Vol/Iss: 11 Pages: 350-351 1955 Author: WHITE et al. Title: Treatment of Pulmonary Hemangiomatosis with Recombinant Interferon Alfa-2a

Examiner	Date	
Signature	Considered	

Vol/lss: 32 (18)

Pages: 1197-1200

Date:

05/04/89

Publ:

The New England Journal of Medicine